Education and debate

"We all have AIDS": case for reducing the cost of HIV drugs to zero

Donald Berwick

An editorial last year in the *Washington Post* challenged directors of drug companies to reduce the cost of HIV drugs in poor countries to zero. Here the reproduced editorial is accompanied by a discussion about why it was written and the response to it. We invited comment from one of the named directors and from a man with AIDS who refuses to take antiretroviral drugs until they are available to the public sector in South Africa

Institute for Healthcare Improvement, 375 Longwood Avenue, Boston, MA 02215, USA Donald M Berwick

president and chief executive officer

dberwick@ihi.org

BMJ 2002;324:214-8

Last year I wrote a guest editorial for the *Washington Post* in which I challenged the world's pharmaceutical companies to cut the cost of HIV drugs to zero in poor countries (see box on next page). Here I explain why I wrote it and describe some of the responses it provoked.

The prologue

People living in poverty being denied access to modern health care is a form of violent, systematic social deprivation that we, as a civilised global community, ought not to accept. Even the poorest people in the poorest settings can, if they are allowed and assisted, be involved in improving their health and can benefit from the most advanced drugs.

This philosophy of social justice drives the international health programme Partners in Health, which was founded by Paul Farmer and Jim Yong Kim. Partners in Health is tackling a seemingly impossible problem: the treatment of multiresistant tuberculosis in a poor shanty town area called Carabayllo on the outskirts of Lima, Peru, and in rural Haiti. The organisation has trained local residents as healthcare "promoters" able to give complex regimens of drugs to patients during directly observed treatment, while providing the patients and their families with a lot of psychosocial support. Success rates have been phenomenal—over 80% of patients have apparently been cured of a disease that only five years ago was thought of in these areas as a death sentence.

Kim's and Farmer's work is changing minds. Until recently, the World Health Organization advised most developing nations not to spend their resources on the diagnostic tools and complex drug regimens used to treat multiresistant tuberculosis in rich nations.³ But now, thanks in part to advocacy from Partners in Health and other organisations, the WHO has at last placed many of the drugs needed to treat multiresistant tuberculosis on its list of essential drugs. Many pharmaceutical manufacturers have reduced the prices of antituberculosis drugs by more than one order of magnitude.⁴

Summary points

Complex drug regimens, combined with social support, have been used to treat infectious diseases, even in conditions of extreme poverty

The high costs of drugs can be used as an excuse for poor countries not to construct effective infrastructures for the care of patients with chronic infectious diseases

Dramatic reductions in the costs of antituberculosis drugs catalysed worldwide action against multiresistant tuberculosis

The boards and executives of drug companies could catalyse action against the AIDS epidemic by immediately reducing the costs of HIV drugs in poor countries to zero

In the international struggle against multiresistant tuberculosis, the sudden and dramatic decrease in the costs of antituberculosis medications was an important catalyst to action. When high costs meant that drugs were far out of reach, it seemed futile for poor countries to try to build infrastructures capable of managing patients with tuberculosis. When drugs became affordable, building a proper healthcare system was a task worth tackling. Peru has begun to broaden the Carabayllo programme to a national scale, and the Bill and Melinda Gates Foundation supports plans to broaden the impact of Partners in Health with multimillion dollar investment.

It takes little imagination to carry one's mind from one scourge—multiresistant tuberculosis—to another—AIDS. ⁵ The challenges are similar, and despair is just as seductive. AIDS is not yet curable, but modern medicine has chipped away at the pace and burden of the disease's progression. Sophisticated drug regimens can increase the healthy portion of the lives of people

with HIV by years, even decades. Worldwide, the stakes are as high as our species has ever known. Without widely available care and effective prevention, AIDS is creating a pandemic without precedent on our planet.⁶ With proper care and active prevention, we could save millions and millions of years of healthy life.^{7 8}

The work of Partners in Health led me, one sleepless night, to write an article about AIDS for an American national newspaper. I wrote a challenge as a guest editorial in the *Washington Post.* Reduce the costs of anti-HIV drugs to zero, or nearly so, I proposed. I directed this challenge at the few people who, with a stroke of their pens, could make it happen the next morning—the executives and boards of the world's pharmaceutical companies.

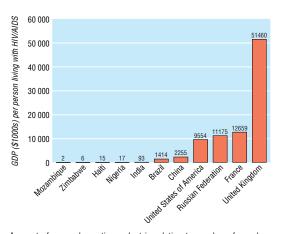
The online debate

The Washington Post hosts question and answer sessions on line on the day that guest editorials are published. I spent two hours answering selected questions from readers. It was an opportunity an author rarely gets—to hear first hand how the reader feels at the moment of reading.

The questions I received provided a cross section of public opinions—some encouraging, some shocking—on our possible role as developed nations in solving the world's AIDS crisis.

Several of the questions debated serious points of fact and evidence.

- Would "attitude, cultural traditions, and gender discrimination" in poor countries "have an adverse effect on the battles against AIDS," even if medications were free? (I replied that removing the barriers of drug costs would, in fact, confront us with the need to tackle such obstacles. High drug costs are an excuse for avoiding other issues.)
- Do "infrastructures" exist to distribute medications and manage treatment? (I cited the success of Partners in Health in developing and sustaining effective infrastructures.)
- Some questions raised concerns about the implications for the pharmaceutical industry if it took my suggestion of making AIDS drugs free.



Amount of gross domestic product in relation to number of people living with HIV infection or AIDS, 1997 (data from www.cptech.org/ip/health/)

The Washington Post editorial

We all have AIDS

In many occupied nations during World War II, the Nazis ordered Jews to wear a yellow star, as prelude to their destruction. But not in Denmark. According to legend, the Danish king, Christian X, threatened that, if Danish Jews were to wear the star, he would, too. The story is almost certainly a myth, but its meaning is not. Despite the Nazi occupation, Denmark rescued the overwhelming majority of its Jews. "If some Danes are under siege," the story means to say, "then all Danes are under siege. So, for now, we are all Jews."

Now we all have AIDS. No other construction is any longer reasonable. The earth has AIDS; 36.1 million people at the end of the year 2000. In Botswana, 36 percent of adults are infected with HIV; in South Africa 20 percent. Three million humans died of AIDS in the year 2000, 2.4 million of them in sub-Saharan Africa. That is a Holocaust every two years; the entire population of Oregon, Iowa, Connecticut or Ireland dead last year, and next year, and next. More deaths since the AIDS epidemic began than in the Black Death of the Middle Ages. It is the most lethal epidemic in recorded history. Prevention will be the most important way to attack AIDS everywhere, but treatment matters, too. We can treat AIDS effectively. We cannot cure its victims, but we can extend their healthy lives by years—with luck, by decades. We can reduce its transmission from infected mother to unborn child by two-thirds or more. We are seeing the effects of advancing science plus enlightened public health policies in the United States, where the toll of AIDS began to fall in 1997.

Successful, life-prolonging management of HIV infection is not simple. Important dimensions include education, social support and life-style interventions that are extremely difficult to achieve in the developed nations, and many times more so in impoverished nations. But it is a mistake to ignore the role of medications. In New York, San Francisco or Nairobi, no matter how different the cultural challenges, the correct mainstay of lifesaving care for the unborn child or the infected adult is medicine, given in a timely, scientifically accurate and reliable way. Most people on earth with HIV and AIDS do not get those medicines. The barriers are partly social and logistical, but the overwhelming barrier is cost. At current prices, one year of triple drug therapy for an HIV-positive person costs \$15,000. Recent, welcome changes by a few progressive pharmaceutical companies, like Merck & Co., promise to reduce that cost by thousands of dollars per year.

But keep in mind that no legend claims King Christian talked of putting on only half a yellow star.

Here is what the world needs: free anti-AIDS medicines. The devastated nations of the world need AIDS medicines at no cost at all, or, at a bare minimum, medicines available at exactly their marginal costs of manufacture, not loaded at all with indirect costs or amortized costs of development. No hand-waving or accounting maneuvers—for all practical purposes, free. Here is how it could happen: the board chairs and executives of the world's leading drug companies decide to do it, period. To the anxious corporate lawyers, the incredulous stockholders, the cynical regulators and the suspicious public, they say, together, the same thing:

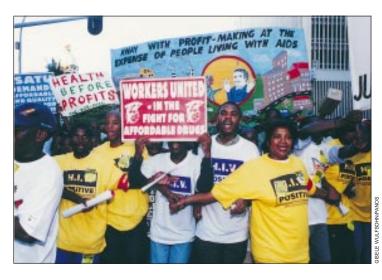
"The earth has AIDS, and therefore we all, for now, have AIDS. Therefore, we are taking one simple action that will save millions and millions of lives. We choose to do it, together, and we will use the intelligence of our own forces to figure out how to make it possible, while preserving the futures of our companies."

No one could stop them; none would dare try. For the small profit they would lose, they would gain the trust and gratitude of the entire world. They would have created a story to be told for a millennium, and those who depend on the prudence of these leaders—on their "fiduciary responsibility"—might chose then not to blame them but to join them in celebration, as fiduciaries of humankind.

The names of the people who can say this, together, include these: Raymond Gilmartin, (chairman and CEO of Merck & Co.); Sir Richard Sykes and Jean-Pierre Garnier (respectively chairman and CEO of GlaxoSmithKline); Charles A Heimbold, Jr, and Peter Dolan (respectively chairman/CEO and president of Bristol-Myers Squibb); Dr Franz B Humer (chairman and CEO of Roche). There are others; they know who they are. These few souls, with this act, would ultimately save the lives of more human beings than died in the Holocaust—perhaps two or three times over.

If a Nobel Prize followed, it would be redundant. The memory of the deed would likely outlive even the story of the Danish king who joined his people in their need.

[C A Heimbold Jr has since retired from Bristol-Myers Squibb.]



- "Where does the r[esearch] and d[evelopment] money come from? Current profits. If you take away the profits, then the companies will have no incentive to do ... research." (I replied that drug companies today get no profits, anyway, from countries that cannot afford their products, and that the international goodwill that could come from bold generosity could put the companies in a much better position to make the case for support for their research agendas.)
- Some readers questioned whether AIDS care deserves such priority.
- "If drug companies ought to provide free AIDS drugs, by the same moral principle they ought to provide free antibiotics and ... every lifesaving drug." (This "Pandora's box" argument is a formula for paralysis. "Let's start somewhere, instead of nowhere," I replied. "And why not with at least one of the greatest scourges we face in the world today?")
- · Questions about where responsibility for dealing with AIDS ought to lie were disturbing to me, because of what they suggested about the potential will for
- "AIDS in most parts of the world is associated with behaviour ... something over which people have some control." (So, should we therefore also not treat other "behaviour induced diseases," such as heart disease, cancer, stroke, and sports injuries?)
- "So who will ultimately end up paying for these drugs? ... [I]t will be just another thing for the tax-

payers to eat? ... How is this [free drugs] supposed to help people help themselves?" ("This is why we have communities," I replied, "including a global community.")

• One email correspondent asked why he should care about AIDS in Africa. "What does this have to do with me?" he asked. "I deeply believe we are one world," I responded, "and all humankind are connected." He replied instantly with a further question, which haunts me still. "Where did you get that idea?" he asked.

Conclusion

Seven months after the editorial appeared, one drug company executive has replied to me. He described and celebrated in a letter important steps his firm has taken to reduce the financial barriers to AIDS care and to support treatment and prevention programmes, steps that I knew about before I wrote the editorial, and that I applaud.

But that is not enough, and it is not what I am asking for. These initial acts of generosity only set the stage for what the world really needs: a dramatic, unprecedented, and unequivocal decision by the boards and executives of several important pharmaceutical companies to make their anti-HIV drugs free. Not half a loaf-a whole loaf. If they did that, these leaders would change the face of the world.

Funding: None.

Competing interests: None declared.

- Berwick DM. We all have AIDS. Washington: Washington Post, Jun 26
- Farmer P, Kim JY. Community based approaches to the control of multidrug resistant tuberculosis: introducing "DOTS-plus." BMJ 1998; 317:671-4.
- World Health Organization. Guidelines for the management of drug-resistant tuberculosis. Geneva: WHO, 1997. Gupta R, Kim JY, Espinal MA, Caudron J-M, Pecoul B, Farmer PE, et al.
- Responding to market failures in tuberculosis control. Science 2001;293:1049-51.
- Farmer P, Léandre F, Mukherjee JS, Claude MS, Nevil P, Smith-Fawzi MC, et al. Community-based approaches to HIV treatment in resource-poor settings. *Lancet* 2001;358:404-9.
- Joint United Nations programme on HIV/AIDS (UNAIDS). AIDS
- epidemic update: December 2000. Geneva: UNAIDS, 2000.

 Mocroft A, Vella S, Benfield TL, Chiesi A, Miller V, Gargalianos P, et al.

 Changing patterns of mortality across Europe in patients with human immunodeficiency virus infection. Lancet 1998;352:1725-30.
- Palella FJ Jr, Delaney KM, Moorman AC, Loveless MO, Fuhrer J, Satten GA, et al. Declining morbidity and mortality among patients with advanced human immunodeficiency virus infection. N Engl J Med 1998:338:853-60.

(Accepted 30 October 2001)

Commentary: The reality of treating HIV and AIDS in poor countries

Richard Sykes

GlaxoSmithKline, Greenford. UB6 0NN Richard Sykes chairman tft28919@

glaxowellcome.co.uk

In the middle of the global AIDS epidemic, it is easy although misguided-to assume that the cost of drugs used to treat HIV and AIDS is the primary barrier to people in poor countries having greater access to such drugs. In reality, the crux of this problem is more fundamental. The main barrier to access is the lack of adequately resourced healthcare systems—as the current British secretary of state for international

development, Clare Short, said, "even if the drugs were free, people would not get them now because there is no delivery system."1

If people are to benefit from antiretroviral treatment, they must have access to counselling and testing facilities, doctors to prescribe appropriate medicines, nurses to teach patients how to take those medicines, and medical aides to ensure that the patients comply with the drug regimen. Sadly, such medical support, and the funding needed to resource it, is rarely available to patients in many poor countries and their governments.

Almost everyone working to improve health care in poor countries agrees that drug donations, although useful in certain circumstances, are not the answer to fighting HIV and AIDS in those countries. The treatment of AIDS must be a lifetime commitment to a patient; therefore, patients and the medical professionals who treat them must be confident that supplies of medicine will be available on a sustainable basis. That is why GlaxoSmithKline offers its medicines to poor countries and countries in sub-Saharan Africa at preferential prices that recover basic costs; this means that the company can make a sustainable commitment to provide its medicines for the long term.

Today, all six of GlaxoSmithKline's HIV and AIDS medicines are available in poor countries at prices up to 90% lower than those charged in rich countries. Despite these price reductions, we have not seen a considerable uptake in the use of these medicines.

So what is the solution? One critical element will be global partnerships created to provide funding and support the development of infrastructures. Kofi Annan, secretary general of the United Nations, estimated the cost of fighting AIDS at \$7bn-\$10bn.² A global health fund would go a long way to supporting the development of infrastructures and prevention programmes, and it would provide a context for effec-

tive distribution and use of drugs. GlaxoSmithKline has already offered to sell its antiretroviral and antimalarial products to the fund at significantly discounted, preferential prices.

The most important part GlaxoSmithKline, and the rest of the pharmaceutical industry, can play is to research and develop new medicines—and to make them available at sustainable preferential prices. GlaxoSmithKline is developing an HIV vaccine, which is perhaps the best hope in the long term for conquering AIDS. We are also working with national governments, agencies of the United Nations, global and African non-governmental organisations, and HIV and AIDS community groups to strengthen the national care and treatment capacities in countries where early results from treatment programmes on small scales are encouraging.

But neither an individual company nor the whole pharmaceutical industry can address this problem on their own. Global partnerships involving all stakeholders—governments, industries, international organisations, non-governmental organisations, community groups interested in HIV and AIDS, the medical profession, and others—are the only way to fight this disease successfully.

Competing interests: RS is chairman of GlaxoSmithKline.

- 1 Clare Short puts pressure on US over AIDS fund. Independent, 23 Jun 2001
- 2 Brown P. Kofi Annan describes new health fund for developing countries. BMJ 2001;322: 1265.

Commentary: Most South Africans cannot afford anti-HIV drugs

Zackie Achmat

If effective action against HIV in South Africa is not taken, between five million and seven million people will have died from AIDS by 2010. Activists have campaigned for more access to antiretroviral drugs to reduce morbidity and mortality related to HIV and AIDS. However, two major obstacles prevent drugs from becoming more widely available—the high prices of the drugs and a lack of political will.

Prices of drugs

In October 2001 the Treatment Action Campaign, which fights for affordable drugs for people infected with HIV in South Africa, organised a meeting of scientists, clinicians, nurses, and civil society organisations. We developed a consensus statement about antiretroviral drugs: "Price reductions should be across the board, and not limited to the public sector. Generic competition of bio-equivalent medicines is essential ... to arrive at the lowest and most sustainable prices for essential medicines."

The brand name oriented portion of the pharmaceutical industry insists that drug prices and patents are not an issue in Africa, or elsewhere. This is not our experience. Currently, Pfizer sells fluconazole in the private sector in South Africa at more than 10 times the price of bioequivalent generic versions sold in Thailand.³ The price that GlaxoSmithKline charges the

public sector for zidovudine is higher than the cost of generic versions manufactured by the Indian company CIPLA and the Brazilian company Farmanguinhos.⁴ We urge all drug companies to issue non-exclusive voluntary licences that allow generic antiretroviral drugs to be produced in and imported into any poor country. A 5% royalty on the generic price should be paid to the patent owners.

Political will

The South African president, Thabo Mbeki, believes that poverty and antiretroviral drugs are the cause of AIDS.⁵ His pseudoscientific beliefs may contribute to the unnecessary death of many poor and black people in South Afrika. Mbeki epitomises leadership in denial, and his stand has fuelled government inaction.

A greater danger exist, however—the international community's lack of political will to provide antiretroviral drugs to people with HIV and AIDS. I believe that the US Agency for International Development, Japanese and European Commission bureaucrats, and British politicians, among others, send out the message that antiretroviral drugs cost too much and that the drug regimens are too complicated for use by people in poor countries. This message is even more cynical and insidious than President Mbeki's lack of political will.

Treatment Action Campaign, Nonkqubela 7793, Cape Town, South Africa Zackie Achmat chair person

Correspondence to: Z Achmat zackie@pixie.co.za

Is infrastructural weakness really the problem?

Sykes says that the weak infrastructure of poor countries will prevent delivery of antiretroviral drugs to those who need it most. This is a poor excuse for inaction. In South Africa, and in other poor countries, many communities would benefit from immediately receiving antiretroviral drugs. Drugs should be distributed while we develop healthcare infrastructures in the poorest communities that are being decimated by HIV and AIDS.

Kofi Annan has asked the G8 countries and other countries to give \$10bn a year towards a global fund for AIDS and health. The global community cannot use this fund to increase the profits of brand name drug companies. It must be used to provide antiretroviral drugs at the lowest possible price. Ten billion dollars will be a start in providing treatment,

supporting HIV prevention efforts, demonstrating political commitment, and reaffirming the principle that everyone has the rights to life, freedom, dignity, and equality.

Competing interests: Zackie Achmat is infected with HIV, but he will not take antiretroviral drugs until they are made available in the public sector in South Africa.

- 1 Dorrington R, Bourne D, Bradshaw D, Laubscher R, Timaeus IM. The impact of HIVAIDS on adult mortality in South Africa. Technical report. Tygerberg: Burden of Disease Research Unit of the Medical Research Council of South Africa, 2001:6.
- 2 Treatment Action Campaign. Bredell consensus statement on the imperative to expand access to anti-retroviral (ART) medicines for adults and children with HIV/AIDS in South Africa. Nonkqubela: Treatment Action Campaign, 2001. www.tac.org.za (accessed 12 Dec 2001).
- 3 Oxfam. Oxfam briefing paper on Pfizer. Oxford: Oxfam, 2001. www.oxfam.org.uk/cutthecost/pfizer.pdf (accessed 12 Dec 2001).
- 4 Yamey G. US trade action threatens Brazilian AIDS programme. BMJ 2001;322:383.
- 5 Sidley P. Clouding the AIDS issue. BMJ 2000;320:1016.

What can we do to reduce mother to child transmission of HIV?

James McIntyre, Glenda Gray

In rich countries antiretroviral treatment in pregnancy has been highly successful in reducing transmission of HIV from mother to child. James McIntyre and Glenda Gray examine whether the success of that treatment—and other strategies, such as changing infant feeding practices—can be repeated in poor countries, which lack both money and infrastructure

Perinatal HIV
Research Unit,
University of the
Witwatersrand,
Chris Hani
Baragwanath
Hospital, PO
Bertsham,
Johannesburg 2013,
South Africa
James McIntyre
director
Glenda Gray
director

Correspondence to: J McIntyre mcintyre@ pixie.co.za

BMJ 2002;324:218-21

HIV infection and AIDS threaten to reverse the gains made in child survival through the more widespread use of childhood vaccines and improved management of diarrhoeal and acute respiratory infections. Although the use of antiretroviral prophylaxis has dramatically reduced mother to child transmission of HIV in the world's rich countries, the effects of these interventions on infant and child survival in poor countries remain undocumented. Our knowledge of the mechanisms and timing of transmission, associated risk factors, and successful methods to reduce the risk have improved over the past decade. Indeed we know how to reduce mother to child transmission of HIV in even the worst affected regions. 1

The World Health Organization has promoted a three pronged approach to reducing mother to child transmission: the prevention of (a) new infections in parents to be, (b) unwanted pregnancies in HIV infected women, and (c) transmission from an HIV infected mother to her infant. The focus to date has been on strategies to prevent transmission to the infant; much is known about these strategies, but the challenge remains in their implementation.

Methods

This article is based on a Medline search of original papers and reviews, unpublished material, presentations and abstracts from recent scientific meetings, information obtained from the internet, and personal communications with scientists and clinicians.

Summary points

Over 600 000 infants worldwide are infected with HIV from their mothers each year

Transmission rates are as high as 35% when there is no intervention and below 5% when antiretroviral treatment and appropriate care are available

Antiretroviral treatment, as long or short course prophylaxis or for treatment of maternal condition, reduces the risk of mother to child transmission

Breast feeding is a major contributing factor to mother to child transmission

Implementing programmes to prevent mother to child transmission has been difficult and slow in poor countries

Future research should focus on prevention of postpartum infection and operational issues

Preventing mother to child transmission

UNAIDS (the joint United Nations programme on HIV and AIDS) estimates that there are over 600 000